UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Box 1450 Alexandria, Virginia 22313-1450 www.uspto.gov

APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.
10/556,485	02/28/2006	Thomas Moritz	2958-136	3791
6449 7590 12/19/2007 ROTHWELL, FIGG, ERNST & MANBECK, P.C. 1425 K STREET, N.W. SUITE 800			EXAMINER	
			MASKELL, MICHAEL P	
WASHINGTON, DC 20005		ART UNIT	PAPER NUMBER	
,			2881	
			NOTIFICATION DATE	DELIVERY MODE
			12/19/2007	ELECTRONIC

# Please find below and/or attached an Office communication concerning this application or proceeding.

The time period for reply, if any, is set in the attached communication.

Notice of the Office communication was sent electronically on above-indicated "Notification Date" to the following e-mail address(es):

PTO-PAT-Email@rfem.com

	Application No.	Applicant(s)				
(						
Office Action Summary	10/556,485	MORITZ ET AL.				
Office Action Summary	Examiner	Art Unit				
The MAILING DATE of this communication app	Michael Maskell	2881				
Period for Reply	ears on the cover sneet with the c	orrespondence address				
A SHORTENED STATUTORY PERIOD FOR REPLY WHICHEVER IS LONGER, FROM THE MAILING DA  - Extensions of time may be available under the provisions of 37 CFR 1.13 after SIX (6) MONTHS from the mailing date of this communication.  - If NO period for reply is specified above, the maximum statutory period w  - Failure to reply within the set or extended period for reply will, by statute, Any reply received by the Office later than three months after the mailing earned patent term adjustment. See 37 CFR 1.704(b).	ATE OF THIS COMMUNICATION 36(a). In no event, however, may a reply be tin will apply and will expire SIX (6) MONTHS from , cause the application to become AB ANDONE	N. nely filed the mailing date of this communication. D (35 U.S.C. § 133).				
Status						
2a) ☐ This action is <b>FINAL</b> . 2b) ☑ This 3) ☐ Since this application is in condition for alloware	Responsive to communication(s) filed on <u>06 September 2006</u> .  This action is <b>FINAL</b> . 2b) This action is non-final.  Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under <i>Ex parte Quayle</i> , 1935 C.D. 11, 453 O.G. 213.					
Disposition of Claims						
<ul> <li>4) ☐ Claim(s) 1-30 is/are pending in the application.</li> <li>4a) Of the above claim(s) is/are withdrawn from consideration.</li> <li>5) ☐ Claim(s) is/are allowed.</li> <li>6) ☐ Claim(s) 1-30 is/are rejected.</li> <li>7) ☐ Claim(s) is/are objected to.</li> <li>8) ☐ Claim(s) are subject to restriction and/or election requirement.</li> </ul>						
Application Papers						
9) The specification is objected to by the Examine 10) The drawing(s) filed on 14 November 2005 is/a Applicant may not request that any objection to the Replacement drawing sheet(s) including the correct 11) The oath or declaration is objected to by the Ex	re: a)⊠ accepted or b)⊡ object drawing(s) be held in abeyance. Se tion is required if the drawing(s) is ob	e 37 CFR 1.85(a). ejected to. See 37 CFR 1.121(d).				
Priority under 35 U.S.C. § 119						
<ul> <li>12) ⊠ Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).</li> <li>a) ⊠ All b) □ Some * c) □ None of:</li> <li>1. ☒ Certified copies of the priority documents have been received.</li> <li>2. □ Certified copies of the priority documents have been received in Application No</li> <li>3. ☒ Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).</li> <li>* See the attached detailed Office action for a list of the certified copies not received.</li> </ul>						
Attachment(s)  1) Notice of References Cited (PTO-892)  2) Notice of Draftsperson's Patent Drawing Review (PTO-948)  3) Information Disclosure Statement(s) (PTO/SB/08) Paper No(s)/Mail Date 11/14/2005.	4) Interview Summary Paper No(s)/Mail D 5) Notice of Informal I 6) Other:	ate				

Application/Control Number: 10/556.485

Art Unit: 2881

#### **DETAILED ACTION**

## Specification

1. The abstract of the disclosure is objected to because it does not include the method of making the claimed article, or the steps in the claimed process. Correction is required. See MPEP § 608.01(b). Applicant is reminded of the proper content of an abstract of the disclosure.

A patent abstract is a concise statement of the technical disclosure of the patent and should include that which is new in the art to which the invention pertains. If the patent is of a basic nature, the entire technical disclosure may be new in the art, and the abstract should be directed to the entire disclosure. If the patent is in the nature of an improvement in an old apparatus, process, product, or composition, the abstract should include the technical disclosure of the improvement. In certain patents, particularly those for compounds and compositions, wherein the process for making and/or the use thereof are not obvious, the abstract should set forth a process for making and/or use thereof. If the new technical disclosure involves modifications or alternatives, the abstract should mention by way of example the preferred modification or alternative.

The abstract should not refer to purported merits or speculative applications of the invention and should not compare the invention with the prior art.

Where applicable, the abstract should include the following:

- (1) if a machine or apparatus, its organization and operation;
- (2) if an article, its method of making;
- (3) if a chemical compound, its identity and use;
- (4) if a mixture, its ingredients;
- (5) if a process, the steps.

Extensive mechanical and design details of apparatus should not be given.

# Claim Rejections - 35 USC § 112

1. The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

2. Claim 12 rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant

10/556,485 Art Unit: 2881

regards as the invention. The claim is apparently incomplete, failing to state what the substrate "moreover comprises." Claim 12 is accordingly not further treated on the merits. Claim 13, being dependent on claim 12, is also therefore indefinite for the same reason and also rejected under 35 U.S.C. 112, second paragraph. Claim 13 is also not further treated on the merits, as it is unknown what limitations claim 13 imports from claim 12.

- 3. Claim 18 rejected under 35 U.S.C. 112, second paragraph, as being indefinite in that it fails to point out what is included or excluded by the claim language. This claim is an omnibus type claim, due to the limitation "providing a sample holder according to the present invention" (emphasis added).
- 4. Claims 18, 19, 26, 28 and 30 rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention. A broad range or limitation together with a narrow range or limitation that falls within the broad range or limitation (in the same claim) is considered indefinite, since the resulting claim does not clearly set forth the metes and bounds of the patent protection desired. See MPEP § 2173.05(c). Note the explanation given by the Board of Patent Appeals and Interferences in *Ex parte Wu*, 10 USPQ2d 2031, 2033 (Bd. Pat. App. & Inter. 1989), as to where broad language is followed by "such as" and then narrow language. The Board stated that this can render a claim indefinite by raising a question or doubt as to whether the feature introduced by such language is (a) merely exemplary of the remainder of the claim, and therefore not required, or (b) a required feature of the claims. Note also, for example, the decisions

10/556,485 Art Unit: 2881

of *Ex parte Steigewald*, 131 USPQ 74 (Bd. App. 1961); *Ex parte Hall*, 83 USPQ 38 (Bd. App. 1948); and *Ex parte Hasche*, 86 USPQ 481 (Bd. App. 1949). In the present instance, claims 18, 26, 28 and 30 recite the broad recitation "phosphorylated/sulphated biopolymers," and the claims also recite "specifically peptides/proteins" which is the narrower statement of the range/limitation. Claim 19 recites the broad limitation "200-600 degrees C" followed by the narrower range "preferably 300 to 450 degrees C" and the even more narrow range "most preferably roughly 400 degrees C," as well as the broad limitation "30 minutes to 180 minutes" followed by the narrower range "preferably 30 to 60 minutes" and the even narrower range "most preferably for roughly 45 minutes."

- 5. Claim 19 rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention. Regarding claim 19, the phrases "for example" renders the claim indefinite because it is unclear whether the limitation(s) following the phrase are part of the claimed invention. See MPEP § 2173.05(d).
- 6. Claims 17 and 19 rejected under 35 U.S.C. 112, second paragraph, as being indefinite for failing to particularly point out and distinctly claim the subject matter which applicant regards as the invention. The term "and/or" in these claims renders the claims indefinite because it is unclear whether the limitations following the term are claimed as required elements. See MPEP § 2173.05(d).

10/556,485 Art Unit: 2881

### Claim Rejections - 35 USC § 102

7. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless -

- (b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.
- 8. Claims 1, 2, 6, 9, 15 and 16 rejected under 35 U.S.C. 102(b) as being anticipated by Tanner, et al (U.S. Patent Application Publication 2003/0003474 A1).

Regarding claim 1, Tanner discloses a sample holder for application in MALDI mass spectrometry (Tanner's invention is a substrate for performing multiple assays of biological materials, and MALDI mass spectrometry is frequently used for assaying biological materials), comprising: a substrate (paragraph 0009); a porous film present on the substrate and comprising metal oxide particles (paragraphs 0009 and 0011).

Regarding claim 2, Tanner discloses the sample holder according to claim 1, characterized in that the metal oxide particles are selected from the group including titanium dioxide, zirconium dioxide, niobium oxide, aluminum titanium oxide, tungsten zirconium oxide, hafnium dioxides, tungsten oxide, tin dioxide, lead oxide, lead dioxide, germanium dioxide, and gallium oxide (paragraph 0011).

Regarding claim 6, Tanner discloses the sample holder according to claim 1, characterized in that the film has a thickness in the range from 0.1 micrometers to 10 micrometers (paragraph 0012).

Regarding claims 7 and 8, Tanner discloses the sample holder according to claim 6, characterized in that the film has a thickness of roughly 3 micrometers

10/556,485 Art Unit: 2881

(paragraph 0012, the terms "roughly" and "about" are interpreted to allow for an up to 50% deviation from the ensuing value; therefore, the thickness of "about" 5 micrometers taught by Tanner indicates a range of 2.5-7.5 micrometers, which overlaps with the "roughly" 3 micrometers range of 1.5-4.5 micrometers).

Regarding claim 9, Tanner discloses the sample holder according to claim 1, characterized in that the substrate consists of glass or coated glass (paragraph 0033).

Regarding claims 15 and 16, Tanner discloses the sample holder according to claim 1, furthermore comprising: one or several samples to be analyzed, which are applied on the film and which is or are presumed to contain one or several substances of interest, selected from the group including nucleic acids and proteins (paragraph 0052).

9. Claims 19, 20 and 24 rejected under 35 U.S.C. 102(b) as being anticipated by Barrow, et al (U.S. Patent 5,585,136).

Regarding claim 19, Barrow discloses a method of preparing a substrate and a porous film applied on the substrate and including metal oxide particles, which method comprises the following steps of operation: preparing a sol from a metal oxide and inducing gel formation, for example by restriction and/or thermal treatment (column 2, line 65-column 3, line 10; Barrow refers to forming the sol gel with ceramic powder, and the definition of "ceramic" according to *The American Heritage*® *Science Dictionary*. Retrieved December 11, 2007, from Dictionary.com website:

<a href="http://dictionary.reference.com/browse/ceramic">http://dictionary.reference.com/browse/ceramic</a> is a material "made typically of metallic

elements combined with oxygen," hence a metal oxide); applying the gel on a substrate,

10/556,485 Art Unit: 2881

drying and subsequent tempering at 200-600 degrees C, preferably 300 to 450 degrees

C, most preferably at roughly 400 degrees C, for a period of 30 minutes to 180 minutes,

preferably 30 minutes to 60 minutes, most preferably for roughly 45 minutes (column 4,

lines 35-49).

Regarding claim 20, Barrow discloses a method according to claim 19, characterized in that the metal oxide particles are selected from the group including titanium dioxide, zirconium dioxide, niobium oxide, aluminum titanium oxide, tungsten zirconium oxide, hafnium dioxides, tungsten oxide, tin dioxide, lead oxide, lead dioxide, germanium dioxide, and gallium oxide (Table 1 lists some of the ceramic powders from which the oxide compounds are formed (column 4, lines 45-50); among these powders are for example Titanium and Zirconium, which would naturally and inherently form

**Regarding claim 24**, this claim is simply the product of the process of claim 19, and is thus anticipated for the same reasons.

titanium dioxide and zirconium oxide, respectively, when oxidized as taught by Barrow).

## Claim Rejections - 35 USC § 103

- 10. The following is a quotation of 35 U.S.C. 103(a) which forms the basis for all obviousness rejections set forth in this Office action:
  - (a) A patent may not be obtained though the invention is not identically disclosed or described as set forth in section 102 of this title, if the differences between the subject matter sought to be patented and the prior art are such that the subject matter as a whole would have been obvious at the time the invention was made to a person having ordinary skill in the art to which said subject matter pertains. Patentability shall not be negatived by the manner in which the invention was made.
- 11. Claims 3-5 rejected under 35 U.S.C. 103(a) as being unpatentable over Tanner in view of Ellson, et al (U.S. Patent Application Publication 2002/0171037 A1). Tanner teaches the sample holder of claim 1, but fails to teach the film having a mean pore size

10/556,485 Art Unit: 2881

of <50 nm, or from 1 nm to 25 nm, or from 1 nm to 10 nm. However, Ellson teaches a porous substrate for holding a MALDI sample with an average pore size of about 10 nm (paragraph 0048). Because these teachings are analogous art (Tanner is directed towards a substrate for multiple biological assays, and Ellson is directed towards analyzing a surface containing a biological sample), one of ordinary skill in the art would be familiar with both teachings; therefore, Ellson's teachings show that a mean pore size of about 10 nm is within the technical grasp of one of ordinary skill in the art. It has been found that "a person of ordinary skill has good reason to pursue the known options within his or her technical grasp. If this leads to the anticipated success, it is likely the product not of innovation but of ordinary skill and common sense." (KSR International Co. v. Teleflex Inc., 550 U.S. , 82 USPQ2d 1385 (2007)). It would therefore have been obvious to one of ordinary skill in the art at the time the invention was made to provide a mean pore size of about 10 nm. Doing so is a known option within the technical grasp of one of ordinary skill, and as taught by Ellson would lead to anticipated success.

12. Claims 10 and 11 rejected under 35 U.S.C. 103(a) as being unpatentable over Tanner in view of Wood, et al (U.S. Patent Application Publication 2004/0094705 A1). Tanner teaches the sample holder according to claim 9, but fails to teach that the glass is a conductive glass or a glass with a conductive coating, or coated with indium tin oxide (ITO). However, Wood teaches a sample holder for MALDI mass spectrometry that may be advantageously used with one or more coatings, one of which is ITO (paragraphs 0095 and 0096). Tanner and Wood's teachings are analogous art because

10/556,485 Art Unit: 2881

Tanner teaches a substrate for biological assays, and Wood teaches a substrate for MALDI mass spectrometry, which is a kind of biological assay; further, Wood teaches a metal oxide coating similar to that taught by Tanner, and teaches that multiple coatings of different types can be used in combination (such as an ITO coating on the glass substrate, followed by the metal oxide film taught by Tanner). Finally, Wood teaches that an ITO coating is advantageous because it provides enhanced desorption, which is desirable for MALDI. It would therefore have been obvious to one of ordinary skill in the art at the time the invention was made to combine the teachings, resulting in the sample holder of claims 10 and 11. Doing so would improve desorption of the sample from the MALDI sample holder.

13. Claim 14 rejected under 35 U.S.C. 103(a) as being unpatentable over Tanner in view of Mayer-Posner, et al (U.S. Patent 6,414,306). Tanner teaches the sample holder according to claim 1, but fails to teach that the film applied on the substrate is present only at defined areas specifically envisaged to this end, covering same, whereas other ranges therebetween are left free of film. However, Mayer-Posner teaches a MALDI mass spectrometry sample carrier plate where the film applied on the substrate is present only at defined areas specifically envisaged to this end, covering same, whereas other ranges therebetween are left free of film (column 8, line 63-column 9, line 8). The teachings are analagous art because Tanner teaches a substrate for performing biological assays and Mayer-Posner teaches MALDI mass spectrometry, which is a kind of biological assay. Further, Mayer-Posner teaches a motivation for the claimed film arrangement, in that samples can be separated yet closely spaced to

increase throughput of analysis. It would therefore have been obvious to one of ordinary skill in the art at the time the invention was made to combine the teachings to provide the sample holder according to claim 14; doing so would gain the throughput advantage taught by Mayer-Posner.

14. Claims 18 and 26-29 rejected under 35 U.S.C. 103(a) as being unpatentable over Tanner in view of Kuroda, et al (U.S. Patent Application Publication 2005/0170525 A1) and in view of Mayer-Posner.

Regarding claim 17, Tanner teaches the sample holder according to claim 16, but fails to teach that the proteins are phosphorylated and/or sulphated. However, Kuroda teaches the analysis of phosphorylated and/or sulphated proteins/peptides using MALDI-MS and a similar metal oxide coating (paragraph 0412). Tanner and Kuroda's teachings are analogous art because Tanner's teachings are directed towards a substrate for biological assays, and Kuroda's teachings are directed towards methods of performing biological assays. Tanner in view of Kuroda makes a strong case for prima facie obviousness for these reasons; however Mayer-Posner is offered to provide further evidence of obviousness. Mayer-Posner teaches a MALDI mass spectrometry sample plate comprising a thin, porous layer containing metal oxide (column 4 lines 64-67). This provides a suggestion in the prior art to combine the substrate coated with a porous metal oxide layer taught by Tanner with the MALDI analysis of phosphorylated peptides taught by Kuroda, by demonstrating that a similar plate has been successfully used in MALDI mass spectrometry. It would therefore have been obvious to one of ordinary skill in the art at the time the invention was made to combine the teachings.

10/556,485 Art Unit: 2881

resulting in the method of claim 17. Doing so is taught to be particularly favorable by Mayer-Posner.

Regarding claim 18, Tanner teaches the sample holder according to claim 1 (this claim is being treated as though "the present invention" refers to claim 1, see 112 indefiniteness rejection above), but fails to teach providing a sample which is presumed to contain phosphorylated/sulphated biopolymers, specifically peptides/proteins, alone or in combination with other biopolymers, specifically peptides/proteins, and applying the sample on the sample holder (the "/" character is being treated as equivalent to the word "or," see 112 indefiniteness rejection above), and performing MALDI mass spectrometry. However, Kuroda teaches providing a sample which contains phosphorylated peptides, applying the sample on a sample holder, and performing MALDI mass spectrometry (paragraph 0412). Tanner and Kuroda's teachings are analogous art because Tanner's teachings are directed towards a substrate for biological assays, and Kuroda's teachings are directed towards methods of performing biological assays. Tanner in view of Kuroda makes a strong case for prima facie obviousness for these reasons; however Mayer-Posner is offered to provide further evidence of obviousness. Mayer-Posner teaches a MALDI mass spectrometry sample plate comprising a thin, porous layer containing metal oxide (column 4 lines 64-67). This provides a suggestion in the prior art to combine the substrate coated with a porous metal oxide layer taught by Tanner with the MALDI analysis of phosphorylated peptides taught by Kuroda, by demonstrating that a similar plate has been successfully used in MALDI mass spectrometry. It would therefore have been obvious to one of

10/556,485 Art Unit: 2881

ordinary skill in the art at the time the invention was made to combine the teachings, resulting in the method of claim 18. Doing so is taught to be particularly favorable by Mayer-Posner.

Regarding claims 26 and 27, as explained in regards to claim 18 above, Tanner teaches the sample holder according to claim 1, but fails to teach the selective detection of phosphorylated/sulfated biopolymers, specifically peptides/proteins, and the detection being performed by means of MALDI mass spectrometry; however, for the same reasons given in regards to claim 18, it would have been obvious to one of ordinary skill in the art at the time the invention was made to combine Tanner with Kuroda in view of Mayer-Posner to apply such a sample and detect it with MALDI mass spectrometry. 15. Claims 21-23 and 24 rejected under 35 U.S.C. 103(a) as being unpatentable over Barrow in view Tanner in view of Ellson. Barrow teaches the method of claim 19 and the sample holder of claim 24, but fails to teach the film having a mean pore size of <50 nm, or from 1 nm to 25 nm, or from 1 nm to 10 nm. Tanner teaches a substrate similar to that produced by Barrow, and teaches its application to biological assays. Ellson teaches a porous substrate for holding a MALDI sample with an average pore size of about 10 nm (paragraph 0048). Because these teachings are analogous art (Barrow is directed towards the method of preparing a substrate, Tanner is directed towards using a similar substrate for multiple biological assays, thus providing the analogous link between Barrow and Ellson, and Ellson is directed towards analyzing a surface containing a biological sample), one of ordinary skill in the art would be familiar with the teachings; therefore, Ellson's teachings show that a mean pore size of about 10

nm is within the technical grasp of one of ordinary skill in the art. It has been found that "a person of ordinary skill has good reason to pursue the known options within his or her technical grasp. If this leads to the anticipated success, it is likely the product not of innovation but of ordinary skill and common sense." (*KSR International Co. v. Teleflex Inc.*, 550 U.S.\_\_\_\_, 82 USPQ2d 1385 (2007)). It would therefore have been obvious to one of ordinary skill in the art at the time the invention was made to provide a mean pore size of about 10 nm. Doing so is a known option within the technical grasp of one of ordinary skill, and as taught by Ellson would lead to anticipated success.

Tanner in view of Kuroda and Mayer-Posner. Tanner teaches providing a sample holder according to claim 1, but fails to teach applying a sample on the metal oxide film of the sample holder, which is presumed to contain phosphorylated/sulphated biopolymers, specifically peptides/proteins; washing the metal oxide film in one or several washing operations; applying a phosphate-containing medium onto the metal oxide film of the sample holder; and applying a MALDI matrix onto the metal oxide film of the sample holder; and the application of this method for performing MALDI mass spectrometry. However, Kuroda teaches applying a sample onto the metal oxide film of the sample holder (paragraph 0409 describes the metal oxide film), which is presumed to contain phosphorylated/sulphated biopolymers, specifically peptides/proteins (paragraph 0412 describes the application and the composition of the sample); washing the metal oxide film in one or several washing operations (paragraph 0411); applying a phosphate-containing medium onto the metal oxide film of the sample holder

10/556,485

Art Unit: 2881

(paragraph 0412); and applying a MALDI matrix onto the metal oxide film of the sample holder and performing MALDI mass spectrometry (paragraph 0142 describes the use of a MALDI plate to detect the peptide via MS; MALDI inherently requires the application of a MALDI matrix, hence the name "Matrix Assisted Laser Desorption Ionization"). These teachings are analogous because Tanner teaches a sample substrate for use in biological assays, and Kuroda teaches the application of such a substrate in a particular biological assay (MALDI-MS). Tanner in view of Kuroda makes a strong case for prima facie obviousness for these reasons; however Mayer-Posner is offered to provide further evidence of obviousness. Mayer-Posner teaches a MALDI mass spectrometry sample plate comprising a thin, porous layer containing metal oxide (column 4 lines 64-67). This provides a suggestion in the prior art to combine the substrate coated with a porous metal oxide layer taught by Tanner with the MALDI analysis of phosphorylated peptides taught by Kuroda, by demonstrating that a similar plate has been successfully used in MALDI mass spectrometry. It would therefore have been obvious to one of ordinary skill in the art at the time the invention was made to combine the teachings, resulting in the method of claims 28 and 29. Doing so would be an application of known elements with no change in their individual function to achieve predictable results.

17. Claim 30 rejected under 35 U.S.C. 103(a) as being unpatentable over Barrow in view of Kuroda and Mayer-Posner. Barrow teaches the sample holder according to claim 24, but fails to teach applying a sample on the metal oxide film of the sample holder, which is presumed to contain phosphorylated/sulphated biopolymers, specifically peptides/proteins; washing the metal oxide film in one or several washing

operations; applying a phosphate-containing medium onto the metal oxide film of the sample holder; and applying a MALDI matrix onto the metal oxide film of the sample holder. However, Kuroda teaches applying a sample onto the metal oxide film of the sample holder (paragraph 0409 describes the metal oxide film), which is presumed to contain phosphorylated/sulphated biopolymers, specifically peptides/proteins (paragraph 0412 describes the application and the composition of the sample); washing the metal oxide film in one or several washing operations (paragraph 0411); applying a phosphate-containing medium onto the metal oxide film of the sample holder (paragraph 0412); and applying a MALDI matrix onto the metal oxide film of the sample holder (paragraph 0142 describes the use of a MALDI plate to detect the peptide via MS; MALDI inherently requires the application of a MALDI matrix, hence the name "Matrix Assisted Laser Desorption Ionization"). These teachings are analogous art because Kuroda teaches the use of a porous metal oxide coating on the MALDI sample substrate, while Barrow teaches the details of forming such a metal oxide coating. Barrow in view of Kuroda makes a strong case for prima facie obviousness for these reasons; however, Mayer-Posner is offered to provide further evidence of obviousness. Mayer-Posner teaches a MALDI mass spectrometry sample plate comprising a thin, porous layer containing metal oxide (column 4 lines 64-67). This provides a suggestion in the prior art to combine the method of forming a substrate coated with a porous metal oxide layer taught by Barrow with the MALDI analysis of phosphorylated peptides taught by Kuroda, by demonstrating that a similar plate has been successfully used in MALDI mass spectrometry. It would have been obvious to one of ordinary skill in the art at the

Art Unit: 2881

time the invention was made to combine the teachings, resulting in the method of claim 30. Doing so is a known application of a substrate analogous to the product produced by Barrow's process, so the combination is an application of known elements together with no change in their individual function.

### Conclusion

18. The prior art made of record and not relied upon is considered pertinent to applicant's disclosure. In particular, the applicant's attention is drawn to Lakshmi, et al (U.S. Patent Application Publication 2004/0248108 A1), which may be available as prior art under 35 U.S.C. 102(e) against the claims as they stand or when amended; but which may be avoided by perfecting applicant's claim to foreign priority by submitting an English translation of the foreign application from which priority is claimed.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Michael Maskell whose telephone number is 571/270-3210. The examiner can normally be reached on Monday-Friday 8AM-5PM EST.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Robert Kim can be reached on 571/272-2293. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see http://pair-direct.uspto.gov. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free). If you would like assistance from a USPTO Customer Service Representative or access to the automated information system, call 800-786-9199 (IN USA OR CANADA) or 571-272-1000.

Michael Maskell

11 December 2007